Please type a plus sign (+) inside this box →

Approved for use through 07/31/96, OMB 0651-0031
Patent and Trademark Office: U.S. DEPARTMENT OF COMMERCE

| | | | | | r accit and Trademark (| JMCE: U.S. DEPARTMENT OF COMMER | CE | |
|---|-------------------------|-------------|-------------|----------------------------------|-------------------------|---------------------------------|----|--|
| | 1449B/PTO Rev. 10/95 | | | nent of Commerce rademark Office | Complete if Known | | | |
| | | | | | Application Number | 09/931,009 | | |
| | LIST O | | | CITED BY | Filing Date | August 17, 2001 | | |
| | | APPI | LICAN" | T | First Named Inventor | Theresa M. SMITH, Ph.D. | | |
| L | (US | e as many s | heets as ne | cessary) | Group Art Unit | 1646 | | |
| Ĺ | | | | | Examiner Name | | | |
| | Sheet | 2 | of | 4 | Attorney Docket Number | US 1257/01 (VA) | | |
| | | • | | - | | | | |

| | ··· | ATIES SPIAN AND AND AND AND AND AND AND AND AND A | |
|-----------------------|--------------|---|---|
| | ···· | OTHER PRIOR ART - NON PATENT LITERATURE DOCUMENTS | |
| Examiner Initials' | Cite No.1 | Include name of the author (in CAPITAL LETTERS), title of the article (when appropriate), title of the item (book), magazine, journal, serial, symposium, catalog, etc.), date, page(s), volume-issue number(s), Publisher, country, where published, source. | 7 |
| 9186 | TO THE | Colvin RB, Johnson RA, Mihm MC, and Dvorak HF. Role of the Clotting System in Cell-Mediated Hypersensitivity. J Exp Med 1973, 138:686-698. | |
| 10V 2 .8 20 | \$ E | Accini L, Dixon FJ. <u>Degenerative Vascular Disease and Myocardial Infarction in Mice with Lupus-like Syndrome</u> . Am J. Pathol 1979, 96:477-492. | |
| & TRADEN | Arr. | Wood RM, Bick MW. The Effect of Heparin on the Ocular Tuberculin Reaction. Arch Opthamol 1959, 61:709-711. | |
| | | Cohen SB, Benacerraf B, McCluskey RT, Ovary Z. Effects of Anticoagulants on Delayed Hypersensitivity Reaction. J Immunol 1967, 98:351-358. | |
| - | | Colvin RB, Dvorak HF. Role of the Clotting System in Cell-Mediated Hypersensitivity II. Kinetics of Fibrinogen/fibrin Accumulation and Vascular Permeability Changes in Tuberculin and Cutaneous Basophil Hypersensitivity Reactions. J Immunol 1975, 114:377-387. | |
| | | Edwards RL, and Rickles FR. <u>Delayed Hypersensitivity in Man: Effects of Systemic Anticoagulation</u> . Science 1978, 200:541-543. | |
| | | Colvin RB, Mosesson MW, Dvorak HF. <u>Delayed Type Hypersensitivity</u> <u>Skin Reactions in Congenital Afibrinogenemia Lack Fibrin Deposition</u> <u>and Induration</u> . J Clin Invest 1979, 63:1302-1306. | |
| | | Malik AB, Johnson A, Tahamont MV. Mechanisms of Lung Vascular Injury After Intravascular Coagulation. Ann NY Acad Sci 1982, 384:213-234. | |
| | | Kay AB, Pepper DS, McKenzie R. The Identification of Fibrinopeptide B as a Chemotactic Agent Derived from Human Fibrinogen. Br J Hematol 1974, 27:669-677. | |
| . | | Richardson DL, Pepper DS, Kay AB. <u>Chemotaxis for Human Monocytes</u> / <u>by Fibrinogen Derived Peptides</u> . Br J Hematol 1976, 32:507-513. | |
| المان | ۲ | Sueishi K, Nanno S, Tanaka K. <u>Permeability Enhancing and Chemotactic Activities of Lower Molecular Weight Degradation Products of Human Fibrinogen</u> . Thromb Haemost 1981, 45:90-94. | |

| Examiner Signature | Sauls | Date Considered | 4-21-04 |
|-----------------------|-------|--------------------|---------|
| | | | <u></u> |

EXAMINER: initial if reference considered, whether or not citation is in conformance with MPEP 609. Draw line through citation if not in conformance and not considered, include copy of this form with next communication to applicant.

¹ Unique citation designation number. ² Applicant is to place a check mark here if English language Translation is attached. Burden Hour Statement: This form is estimated to take .2 hours to complete. Time will vary depending upon the needs of the individual case. Any comments on the amount of time you are required to complete this form should be sent to the Chief Information Officer, Patent and Trademark Office, Washington, DC 20231. DO NOT SEND FEES OR COMPLETED FORMS TO THIS ADDRESS. SEND TO: Assistant Commissioner for Patents.



PTO/SB/08B (6-95)
Approved for use through 07/31/96, OMB 0651-0031

| Please to | pe a plus sign (+) ins | ide this box | → | Approved for use through 07/31/96, OMB 0651-0031 Patent and Trademark Office: U.S. DEPARTMENT OF COMMERCE | | | |
|-----------------------|------------------------|--------------|-----------------------------------|--|-------------------------|--|--|
| 1449B/P' Rev. 10/9 | | | ent of Commerce ademark Office | Complete if Known | | | |
| | | | | Application Number | 09/931,009 | | |
| LIST | OF PRIOR | R ART | CITED BY | Filing Date | August 17, 2001 | | |
| | APPL | ICANT | • | First Named Inventor | Theresa M. SMITH, Ph.D. | | |
| | (use as many sh | eets as nec | essary) | Group Art Unit | 1646 | | |
| | | | | Examiner Name | | | |
| Sheet | 3 | of | 4 | Attorney Docket Number | US 1257/01 (VA) | | |

| OTHER PRIOR ART - NON PATENT LITERATURE DOCUMENTS | | | | | | | |
|---|-------------------|--|--------------|---|----|--|--|
| | Exami Initials | | Cite No.1 | Include name of the author (in CAPITAL LETTERS), title of the article (when appropriate), title of the Item (book), magazine, journal, serial, symposium, catalog, etc.), date, page(s), volume-issue number(s), Publisher, country, where published, source. | T² | | |
| | Su 61 | | - 20 | Rowland F, Donovan M, Gillies C, O'Rourke J, Kreutzer DL. <u>Fibrin:</u> <u>Mediator of In Vivo and In Vitro Injury and Inflammation</u> . Curr Eye Res 1985, 4:537-553. | | | |
| વ્ર | NON 1 | | See See | Saxne T, Lecander I, Geborek P. <u>Plasminogen Activators and Plasminogen Activator</u> <u>Inhibitors in Synovial Fluid Difference Between Inflammatory Joint Disorders and Osteoarthritis</u> . J Rheumatol 1993, 20:91-96. | | | |
| | Ta T | RADE | MARK | Kikuchi H, Tanaka S, Matsuo O. <u>Plasminogen Activator in Synovial Fluid from Patients with Rheumatoid Arthritis</u> . J Rheumatol 1987, 14:439-445. | | | |
| | | Kummer JA, Abbink JJ, De Boer JP, Roem D, Nieuwenhuys EJ, Kamp AM, Swa Hack CE. Analysis of Intraarticular Fibrinolytic Pathways in Patients with Inflammand Noninflammatory Joint Diseases. Arthritis Rheum 1992, 35:884-893. | | | | | |
| | | Belch JJF, McArdle B, Madhok R. <u>Decreased Plasma Fibrinolysis in Patients with Rheumatoid Arthritis</u> . Ann Rheum Dis 1984, 43:774-77 | | | | | |
| | | | , | Dahlquist SR, Jonsson SW, Ranby M. <u>Fibrinolytic Components in Synovial Fluid of Destructive and Non-Destructive Arthritis</u> . Arthritis Rheum 1994, 37:S248. | | | |
| | | | | Herrick AL, Illingworth K, Blann A, Hollis S, Jayson MIV. <u>Von Willebrand</u> <u>Factor, Thromboxane, B-Thromboglobulin and Markers of Fibrinolysis in Primary</u> <u>Raynaud's Phenomenon and Systemic Sclerosis</u> . Arthritis Rheum 1994, 37:S259. | | | |
| | | | | Qi J, Kreutzer DL, Piela-Smith TH. <u>Fibrin Induction of ICAM-1 Expression in Human Vascular Endothelial Cells</u> . J Immunol 1997, 158:1880-1886. | | | |
| | | | | Kawasaki K, Hirase K, Masanori M, Tsuji T, Iwamoto, M. <u>Amino Acids and Peptides XVI. Synthesis of N-terminal Tetrapeptide Analogs of Fibrin a Chain and Their Inhibitory Effects on Fibronogen/fibrin Clotting.</u> 1992. Chem Pharm Bull 40:3253-3260. | - | | |
| | | | | Cliffton EE, Grossi CE. <u>The Rationale of Anticoagulants in the Treatment of Cancer</u> . J Med 1974, 5:107-113. | | | |
| L | Sir | ١٤ | | Peterson HI. Fibrinolysis and Antifibrinolytic Drugs in the Growth and Spread of Tumors. Cancer Treat Rev 1977, 4:213-217. | | | |

| Examiner Signature Date Considered Considered | | |
|---|-----|--------------------|
| | SMS | Q-20-U4/1-3105 |

EXAMINER: Initial if reference considered, whether or not citation is in conformance with MPEP 609. Draw line through citation if not in conformance and not considered. Include copy of this form with next communication to applicant.

¹ Unique citation designation number. ² Applicant is to place a check mark here if English language Translation is attached.

Burden Hour Statement: This form is estimated to take .2 hours to complete. Time will vary depending upon the needs of the individual case. Any comments on the amount of time you are required to complete this form should be sent to the Chief Information Officer, Patent and Trademark Office, Washington, DC 20231. DO NOT SEND FEES OR COMPLETED FORMS TO THIS ADDRESS. SEND TO: Assistant Commissioner for Patents, Washington, DC 20231.

| PTO/SB/08 | B (6-95) |
|---|----------|
| Approved for use through 07/31/96, OMB 0 | 551-0031 |
| Patent and Trademark Office: U.S. DEPARTMENT OF COM | MERCE |

| · | 1 10000 1700 1 | , prop 5-m-1 () | | | Patent and Trademark C | Office: U.S. DEPARTMENT OF COMMERC | Œ |
|---|-------------------------|------------------|----------------------------------|-----------------------------|------------------------|------------------------------------|---|
| (| 14498/PTO Rev. 10/95 | | S. Department atent and Trade | of Commerce emark Office | C | omplete if Known | |
| | | | | | Application Number | 09/931,009 | |
| ĺ | LIST C | F PRIO | R ART C | ITED BY | Filing Date | August 17, 2001 | |
| | | APPI | LICANT | | First Named Inventor | Theresa M. SMITH, Ph.D. | |
| | (0 | ise as many s | heets as neces | ssary) | Group Art Unit | 1646 | |
| | | | | | Examiner Name | | |
| | Sheet | 4 | of | 4 | Attorney Docket Number | US 1257/01 (VA) | |

| | ATUED DDIAD ADT NAMBATENT LITERATURE DAGULLENTA | |
|------------------------------|--|----------|
| | OTHER PRIOR ART - NON PATENT LITERATURE DOCUMENTS Include name of the author (in CAPITAL LETTERS), title of the article (when appropriate), title of the | \vdash |
| Examiner Cite Initials PNo | Item (book), magazine, journal, serial, symposium, catalog, etc.), date, page(s), volume-issue number(s), Publisher, country, where published, source. | T² |
| MOV 2 8 2001 | Cacharski LR, Henderson WG, Rickles FR, Forman WB, Cornell CJ Jr, Forcier RJ, Sowards R, Headley E, Kim SH, O'Donnell JR, O'Dell R, Tornyos K, Kwaan HC. Effect of Warfarin on Survival in Small Cell Carcinoma of the Lung. JAMA 1981, 245:831-835. | |
| TA ADENIUM | Bardos H, Juhasz A, Repassy G, Adany R. <u>Fibrin Deposition in Squamous</u> Cell Carcinomas of the Larynx and Hypopharynx. Thromb Haemost 1998, 80:767-772. | |
| | Laudano AP, Doolittle RF. Studies on Synthetic Peptides That Bind to Fibrinogen and Prevent Fibrin Polymerization. Structural Requirements, Number of Binding Sites, and Species Differences. 1980, Biochemistry 19:1013-1019. | |
| | Shreck R, Meier B, Manne D, Droge W, Bauerle PA. <u>Dithiocarbamate as Potent Inhibitors of Nuclear Factor kB in Intact Cells</u> . J Exp Med 1992, | |
| | Lorenz H-M, Kalden JR. <u>Biologic Agents in the Treatment of Inflammatory</u> Rheumatic Diseases. 1999, Curr Opin Rheumatol 11:179-184. | |
| | Rothchild, BM, Thompson, LD, Pifer DD, Chesney CM. Perturbation of Protease Inhibitors and Substrates in Inflammatory Arthritis. Semin Thromb Hemostasis 1985, 11:394-404. | ; |
| SWL | Belch JJF, Madhok R, McArdle B, McLaughlin K, Kluft C, Forbes CD, Sturrock RD. The Effect of Increasing Fibrinolysis in Patients with Rheumatoid Arthritis: A Double Blind Study of Stanozolol. Q J Med 1986, 58:19-27. | |
| | | |
| | | |
| | | |
| | | |

| 4 | | | | |
|---|-----------------------|------|--------------------|----------------|
| | Examiner Signature | Suts | Date Considered | 4-20-4/1-31-05 |
| | | | | |

EXAMINER: Initial if reference considered, whether or not citation is in conformance with MPEP 609. Draw line through citation if not in conformance and not considered. Include copy of this form with next communication to applicant.

¹ Unique citation designation number. ² Applicant is to place a check mark here if English language Translation is attached.

Burden Hour Statement: This form is estimated to take .2 hours to complete. Time will vary depending upon the needs of the individual case. Any comments on the amount of time you are required to complete this form should be sent to the Chief Information Officer, Patent and Trademark Office, Washington, DC 20231. DO NOT SEND FEES OR COMPLETED FORMS TO THIS ADDRESS. SEND TO: Assistant Commissioner for Patents, Washington, DC 20231.

PTO/SB/08A (6-95)

| Please type a plu | s sign (+) îns id i | e this box → | | Approved for use through 9/30/98. OMB 0651-0031 Patent and Trademark Office: U.S. DEPARTMENT OF COMMERCE | | |
|-------------------------|----------------------------|--------------|----------------------------|--|-------------------------|--|
| 1449B/PTO Rev. 10/95 | | | of Commerce mark Office | C mplete if Known | | |
| | | | | Application Number | 09/931,009 | |
| LIST OF | PRIOR | ART CI | TED BY | Filing Date | August 17, 2001 | |
| | APPLI | CANT | | First Named Inventor | Theresa M. SMITH, Ph.D. | |
| (use e | s many shee | ts as neces | sary) | Group Art Unit | 1646 | |
| | | | | Examiner Name | | |
| Sheet | 1 | of | 4 | Attorney Docket Number | US 1257/01 (VA) | |

| U.S. PATENT DOCUMENTS | | | | | | | | |
|-----------------------|------|--------------|--|--|---|--------------|--|--|
| Examiner Initials' | | | Name of Patentee or Applicant of Cited Document | Date of Publication of Cited Document MM-DD-YYYY | Pages, Columns, Lines, Where Refevant Passages or Refevant Foures Annear | | | |
| | A | 3410 | | | | | | |
| | / | <u> </u> | \ _ | | | | | |
| | 1- N | V 7 H 2000 E | 1 | | | V- | | |
| | 13 | · ¿ o zun | ,} | | (J ¹) | | | |
| | 135 | | | | | 107 | | |
| | -1 | W Z B ZOON E | | | | \ | | |
| | | | | | | 1,1,1 | | |
| | | | | (| , , | | | |
| | | | | 0.0 | | | | |
| | | | | | | | | |
| | | | | | | | | |
| <u> </u> | | · | | | 1 | | | |
| | | | | D. / | | | | |
| | | | +{ | | | | | |
| | | ···· | | | | | | |
| | | | | | | | | |
| | | | | | | | | |
| | | | | | | | | |

| · | | | | | POREIG | N PATENT DOCUMENT | S | | - |
|-----------------------|--------------|-------------------------------------|--|--|--------|--|---------------------------|--|--|
| Examiner Initials' | Cite No.1 | Foreign Patent Document | | | | No. of Board | Date of Publication of | Pages, Columns, Lines, | T |
| | | Office Number Kind Codes (if known) | | | | Name of Patentee or Applicant of Cited Document | Cited Document MM-DD-YYYY | Where Relevant Passages or Relevant Figures Appear | To |
| | | | | | | | | | |
| | | <u> </u> | | | | | | | 1 |
| | <u> </u> | | | | | | | | |
| · | | | | | | | | | \vdash |
| | L | | | | | | | | ├── |
| | | | | | | | <u> </u> | | ├ |
| | | | | | | ··· | | | ├ |
| | / | | | | | | | | ├ |
| | | | | | | | | | ├ |
| | | | | | | | | | |

| Examiner Signature | | Date Considered | | | | | | | |
|-----------------------|--|--------------------|--|--|--|--|--|--|--|
| | | | | | | | | | |

EXAMINER: Initial if reference considered, whether or not citation is in conformance with MPEP 609. Draw line through citation if not in conformance and not considered. Include copy of this form with next communication to applicant.

¹ Unique citation designation number. ² See attached Kinds of U.S. Patent Documents. ³ Enter Office that issued the document, by the two-letter code (WIPO Standard ST.3) ⁴ For Japanese patent documents, the indication of the year of the reign of the Emperor must precede the serial number of the patent document. ⁵ Kind of document by the appropriate symbols as indicated on the document under WIPO Standard ST.1 ⁶ if possibile. ⁶ Applicant is to place a check mark here if English language Translation is attached.



Burden Hour Statement: This form is estimated to take .2 hours to complete. Time will vary depending upon the needs of the individual case. Any comments on the amount of time you are required to complete this form should be sent to the Chief Information Officer, Patent and Trademark Office, Washington, DC 20231. Under the Paperwork Reduction Act of 1995, no persons are required to respond to a collection of Information unless it displays a valid OMB control number. DO NOT SEND FEES OR COMPLETED FORMS TO THIS ADDRESS. SEND TO: Assistant Commissioner for Patents, Washington, DC 20231.